

GenCore version 5.1.3  
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## Om protein - protein search, using sw model

Run on: November 30, 2002, 12:31:03 ; Search time 27 Seconds

(without alignments)  
2482.410 Million cell updates/secTitle: US-10-025-514-16  
Perfect score: 2675

Sequence: 1 MEDPOGDAAQKTDTSHHDD. ....RDJLKCNCMGKSCVSPVKA 503

Scoring table: BLOSUM62

Gapop 10.0 , Gapext 0.5

Searched: 908470 seqs, 133250620 residues

Total number of hits satisfying chosen parameters: 908470  
Minimum DB seq length: 0  
Maximum DB seq length: 2000000000Post-processing: Minimum Match 0%  
Maximum Match 100%

Listing first 45 summaries

Database : A\_Geneseq\_101002:\*

1: /SIDS2/gcdata/geneseq/geneseqp-emb1/AA1980.DAT:\*

2: /SIDS2/gcdata/geneseq/geneseqp-emb1/AA1981.DAT:\*

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22: /SIDS2/gcdata/geneseq/geneseqp-emb1/AA2001.DAT:\*

23: /SIDS2/gcdata/geneseq/geneseqp-emb1/AA2002.DAT:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## SUMMARIES

Result No.	Score	Query Match	Length	DB ID	Description
1	2675	100.0	503	23 AAU99884	rSLAP1 fusion prot
2	240.5	76.3	522	23 AAU99885	rN-TAP1 fusion pro
3	240.5	76.3	580	23 AAU99889	rTAP1 fusion prote
4	2035	76.1	503	23 AAU99881	NrTAP1 fusion prote
5	2035	76.1	522	23 AAU99883	TAP1 fusion prote
6	2035	76.1	580	23 AAU99882	Mature Protein seq
7	2030	75.9	394	19 AAU99839	Human alpha-1-anti
8	2030	75.9	394	23 AAU99873	Sequence of human AAT
9	2030	75.9	418	5 AAP40133	Predominant form o
10	2030	75.9	418	10 AAP54664	

## SUMMARIES

Human alpha-1-antitrypsin

Alpha-1-antitrypsin

Human alpha-1-tryptase

Amino acid sequence

Human alpha-1-antit

Human alpha-1-protease

Human alpha-1-antit

Human alpha-1-antit

Sequence encoded b

[Leu358] alpha-1-an

Entire sequence of

[Ile358] alpha-1-an

[Ile358] alpha-1-an

Sequence of alpha-

Alpha-1 antitrypsin

[Phe358] alpha-1-an

Alpha-1-antitrypsin

[Ala358] alpha-1-an

[Arg358] alpha-1-an

Sequence of alpha-

[Gly358] alpha-1-an

Sequence encoded b

[Phe358] alpha-1-an

Alpha-1-antitrypsin

[Ala358] alpha-1-an

[Arg358] alpha-1-an

Sequence of alpha-

[Gly358] alpha-1-an

Sequence encoded b

[Ala358] alpha-1-an

Alpha-1-antitrypsin

[Ala358] alpha-1-an

Alpha-1-antitrypsin

[Ala358] alpha-1-an

Human alpha-1-antit

Sequence of human

[Gly358] alpha-1-an

Sequence of human

[Ala358] alpha-1-an

PF	18-DEC-2001;	2001WO-US49256.	Qy	421	ECQSDWQCPGKKRCCPDTGGIKCLDPVDTPNPTREKGKCPVTYGOCLMLNPPNFCMDG 480
XX			Db	421	ECQSDWQCPGKKRCCPDTGGIKCLDPVDTPNPTREKGKCPVTYGOCLMLNPPNFCMDG 480
PR	18-DEC-2001;	2000US-256599P.	Qy	481	QCKRDILKCMGKSCVSPVKA 503
PR	20-NOV-2001;	2001US-331966P.	Db	481	QCKRDILKCMGKSCVSPVKA 503
PA	(ARRI-)	ARRIVA PHARM INC.			
XX					
PI	Barr PJ,	Gibson HL,	Pemberton P;		
XX					
DR	WPI: 2002-500631/53.		RESULTS 2		
DR	N-PSDB;	ABK88025.	ID	AAD99885	standard; Protein: 522 AA.
XX			XX	AAD99885;	
PT	Novel fusion protein useful for inhibiting protease activity associated with a disorder such as emphysema, asthma, comprises a first protease		AC	AAD99885;	
PT	inhibitor comprising alpha 1-antitrypsin and a second protease		XX		
PT	inhibitor -		DT	07-OCT-2002	(first entry)
XX			XX		
BS	Example 3; Page 90-91; 134pp; English.		DE		rN-TAPI fusion protein.
XX	This invention relates to a novel fusion protein comprising a first		XX		rn-TAPI; Alzheimer's disease; tumour angiogenesis;
CC	protease inhibitor comprising an alpha 1-antitrypsin or its functionally		XX		KW malaria; emphysema; asthma; chronic obstructive pulmonary disease;
CC	active portion and a second protease inhibitor or its functionally		XX		KW cystic fibrosis; otitis media; otitis external; HIV; psoriasis; eczema;
CC	active protein. The fusion proteins of the invention may act as an		XX		KW human immunodeficiency virus; atopic dermatitis; muscular dystrophy;
CC	inhibitor of protease activity. The fusion protein of the invention		XX		KW herpes; ulceration; sepsis; rheumatoid arthritis; periodontal disease;
CC	is useful for inhibiting protease activity associated with a disorder		XX		KW tumour metastasis; osteoporosis; scleroderma; Paget's disease; hypertension;
CC	such as emphysema, asthma, chronic obstructive pulmonary disease,		XX		KW glomerulonephritis; hypertension;
CC	cystic fibrosis; otitis media; otitis external or HIV infection, or		XX		KW Homo sapiens.
CC	for treating an individual suffering from or at risk for a disease or		OS		OS Synthetic.
CC	disorder involving unwanted protease activity. The proteins are useful		XX		
CC	for treating dermatological diseases such as atopic dermatitis, eczema		FH		
CC	and psoriasis, in inflammatory responses to viral infection, and for		Region	2..395	Location/Qualifiers
CC	treating herpes infection, corneal or epidermal ulceration, chronic		FT		"Human AAT amino acids 1-394"
CC	non-healing wounds, sepsis, rheumatoid arthritis, periodontal disease,		FT		/note= "Linker methionine"
CC	tumour metastasis and tumour angiogenesis, gastric ulceration,		FT	396	
CC	osteoporosis, Paget's disease, glomerulonephritis, scleroderma, malaria,		FT	397..522	
CC	bacterial infection, Alzheimer's disease, hypertension and muscular		FT		/note= "Amino acids 1-126 of human TIMP-1"
CC	dystrophy. The present sequence represents the rN-TAPI fusion protein of		XX		
CC	the invention.		XX		WO200250287-A2.
XX			XX		
SQ	Sequence 503 AA;		XX		
Query	Match 100.0%; Score 2675; DB 23; Length 503;		XX		
Best Local Similarity 100.0%; Pred. No. 1e-199;			PD	27-JUN-2002.	
Matches 503; Conservative 0; Mismatches 0; Indels 0; Gaps 0;			XX		
Db	1 MEQDQGDAAQKDTSHDQDPTFNKTPNAAEFAFSLYQLQLAHSNSTNFFSPVSIAT 60		PF	18-DEC-2001; 2001WO-US49256.	
Db	1 MEQDQGDAAQKDTSHDQDPTFNKTPNAAEFAFSLYQLQLAHSNSTNFFSPVSIAT 60		XX		
Qy	1 AFAMLSLGTKAETHDELEGIFNLFTEIPAEQTHFGQELLTNLQDPSQQLTGNGLF 120		PR	18-DEC-2000; 2000US-256599P.	
Db	61 AFAMLSLGTKAETHDELEGIFNLFTEIPAEQTHFGQELLTNLQDPSQQLTGNGLF 120		XX	20-NOV-2001; 2001US-331966P.	
Qy	121 LS8GLKLVDKFLEDVKLHYSEAFTYNGDPEAKQINDYVEKGTQKLVYKELDRD 180		PA	(ARRI-) ARRIVA PHARM INC.	
Db	121 LS8GLKLVDKFLEDVKLHYSEAFTYNGDPEAKQINDYVEKGTQKLVYKELDRD 180		XX		
Qy	181 TPLAVNTTFRKWERFEWDEEDFDDQTVTVKVPARKLGMENLQHCKLSSWV 240		PI	Barr PJ, Gibson HL, Pemberton P;	
Db	181 TPLAVNTTFRKWERFEWDEEDFDDQTVTVKVPARKLGMENLQHCKLSSWV 240		XX		
Qy	241 LLMKYLNATAIIFLPDGEKLOHLENELTHDITKFDENEDERSASLHLPLSITGTYDL 300		DR	2002-500631/53.	
Db	241 LLMKYLNATAIIFLPDGEKLOHLENELTHDITKFDENEDERSASLHLPLSITGTYDL 300		DR	N-PSDB; ABK88027.	
Qy	241 KSVLGQLGTTKVFSNGADLSGYTEAPKLSSKAVHKAYLTDEKGTEAGAMFLEATPMS 360		PS	Example 3; Page 97; 134pp; English.	
Db	301 KSVLGQLGTTKVFSNGADLSGYTEAPKLSSKAVHKAYLTDEKGTEAGAMFLEATPMS 360		XX	This invention relates to a novel fusion protein comprising a first	
Qy	301 KSVLGQLGTTKVFSNGADLSGYTEAPKLSSKAVHKAYLTDEKGTEAGAMFLEATPMS 360		CC	protease inhibitor comprising an alpha 1-antitrypsin or its functionally	
Db	361 IPPEVKPNKPFPVFLMEQNTKSPLLEMGVNVNPTQKMSGSFKSFAGYCPPKSAOCLRYKKP 420		CC	active portion and a second protease inhibitor or its functionally	
Qy	361 IPPEVKPNKPFPVFLMEQNTKSPLLEMGVNVNPTQKMSGSFKSFAGYCPPKSAOCLRYKKP 420		CC	active protein. The fusion proteins of the invention may act as an	
Db	361 IPPEVKPNKPFPVFLMEQNTKSPLLEMGVNVNPTQKMSGSFKSFAGYCPPKSAOCLRYKKP 420		CC	inhibitor of protease activity. The fusion protein of the invention	
			CC	is useful for inhibiting protease activity associated with a disorder	
			CC	such as emphysema, asthma, chronic obstructive pulmonary disease,	
			CC	cystic fibrosis, otitis media, otitis external or HIV infection, or	
			CC	disorder involving unwanted protease activity. The proteins are useful	
			CC	for treating an individual suffering from or at risk for a disease or	
			CC	disorder involving dermatological diseases such as atopic dermatitis, eczema	
			CC	and psoriasis, in inflammatory responses to viral infection, and for	

treating herpes infection, corneal or epidermal ulceration, chronic non healing wounds, sepsis, rheumatoid arthritis, periodontal disease, tumour metastasis and tumour angiogenesis, gastric ulceration, osteoporosis, Paget's disease, glomerulonephritis, scleroderma, malaria, bacterial infection, Alzheimer's disease, hypertension and muscular dystrophy. The present sequence represents the rN-rAPI fusion protein of the invention.

Sequence 522 AA:

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Query Match 76.3%; Score 2040.5; DB 23; Length 522;
Best Local Similarity 96.6%; Pred. No. 2, 7e-150;
Matches 400; Conservative 2; Mismatches 7; Indels 5; Gaps 1;
Qy 1 MEDPQGDAAQKTDPTSHHDQDHPTENKTPNLAEAFASLYROLAQSNTNFESPVSTAT 60
Db 1 MEDPQGDAAQKTDPTSHHDQDHPTENKTPNLAEAFASLYROLAQSNTNFESPVSTAT 60
Qy 61 AFAMLSLGTKAQTHDETLGLANFLNLTPEAQIHEGFOELRTLNOPDSQLQTONGLF 120
Db 61 AFAMLSLGTKAQTHDETLGLANFLNLTPEAQIHEGFOELRTLNOPDSQLQTONGLF 120
Qy 121 LSEGKLKVLDKFLEDVKLYHSRAFTVNFGDTEAKQKINDVYKGQTKIVDVLKELDRD 180
Db 121 LSEGKLKVLDKFLEDVKLYHSRAFTVNFGDTEAKQKINDVYKGQTKIVDVLKELDRD 180
Qy 181 TYPALVNYYFFKGKWERPFEVKDTEERDFHYDQVTTVKVPMKMRGLMENIOHCKKLSSWY 240
Db 181 TYPALVNYYFFKGKWERPFEVKDTEERDFHYDQVTTVKVPMKMRGLMENIOHCKKLSSWY 240
Qy 241 LLMKYLGNATAIFFLPDEGKQLOHLENLTDITKFLENEDRRSASILHPLKSITGTYDL 300
Db 241 LLMKYLGNATAIFFLPDEGKQLOHLENLTDITKFLENEDRRSASILHPLKSITGTYDL 300
Qy 301 KSVLGQLGCTKVFNSNGADLSGYTEEAPLKLSKAVHAVITIDEKGTEAGAMFLERIPMS 360
Db 301 KSVLGQLGCTKVFNSNGADLSGYTEEAPLKLSKAVHAVITIDEKGTEAGAMFLERIPMS 360
Qy 361 IPEBVKPNKPFPVFLMBQNTKSPLFMGVNVNPTQKMSKSFKAQGCPPKSAQC 414
Db 361 IPEBVKPNKPFPVFLMBQNTKSPLFMGVNVNPTQKMSKSFKAQGCPPKSAQC 409

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## RESULT 3

AAU99889  
ID AAU99889 standard; Protein: 580 AA.

AC AAU99889;

XX DT 07-OCT-2002 (first entry)

XX DE rTAPl fusion protein.

XX KW rTAPl; Alzheimer's disease; tumour angiogenesis; malaria; emphysema; asthma; chronic obstructive pulmonary disease; cystic fibrosis; otitis media; otitis externa; HIV; poriasis; eczema; human immunodeficiency virus; atopic dermatitis; muscular dystrophy; herpes; ulceration; sepsis; rheumatoid arthritis; periodontal disease; tumour metastasis; osteoporosis; Paget's disease; scleroderma; glomerulonephritis; hypertension.

XX OS Homo sapiens.  
OS Synthetic.

XX FH Key

FT Region 2..395  
/note= "Human AAT amino acids 1-394"

FT Region 396  
/note= "Linker methionine"

FT Region 397..580  
/note= "Amino acids 1-184 of human TIMP-1"

XX PN WO200250287-A2.

XX PD 27-JUN-2002.
XX PF 18-DEC-2001; 2001WO-US49256.
XX PR 18-DEC-2000; 2000US-25669P.
XX PR 20-NOV-2001; 2001US-331966P.
XX PA (ARRI) ARRIVA PHARM INC.
XX PT Barr PJ, Gibson HL, Pemberton P;
XX DR WPI; 2002-500631/53.
XX PT Novel fusion protein useful for inhibiting protease activity associated with a disorder such as emphysema, asthma, comprises a first protease inhibitor comprising alpha 1-antitrypsin and a second protease inhibitor -
XX PT Example 3; Page 94; 134PP; English.
XX PS Example 3; Page 94; 134PP; English.
XX CC This invention relates to a novel fusion protein comprising a first protease inhibitor comprising an alpha1-antitrypsin or its functional active portion and a second protease inhibitor or its functional active protein. The fusion proteins of the invention may act as an inhibitor of protease activity. The fusion protein of the invention is useful for inhibiting protease activity associated with a disorder such as emphysema, asthma, chronic obstructive pulmonary disease, cystic fibrosis, otitis media, otitis externa or HIV infection, or for treating an individual suffering from or at risk for a disease or disorder involving unwanted protease activity. The proteins are useful for treating dermatological diseases such as atopic dermatitis, eczema and psoriasis, in inflammatory responses to viral infection, and for treating herpes infection, corneal or epidermal ulceration, chronic non-healing wounds, sepsis, rheumatoid arthritis, periodontal disease, tumour metastasis and tumour angiogenesis, gastric ulceration, osteoporosis, Paget's disease, glomerulonephritis, scleroderma, bacterial infection, Alzheimer's disease, hypertension and muscular dystrophy. The present sequence represents the rTAPl fusion protein of the invention.
XX SQ sequence 580 AA;
XX Query Match 76.3%; Score 2040.5; DB 23; Length 580;
Best Local Similarity 96.6%; Pred. No. 3.1e-150;
Matches 400; Conservatory 2; Mismatches 7; Indels 5; Gaps 1;
Db 1 MEDPOGDAAQKTDPTSHHDQDHPTENKTPNLAEAFASLYROLAQSNTNFESPVSTAT 60
Db 1 MEDPOGDAAQKTDPTSHHDQDHPTENKTPNLAEAFASLYROLAQSNTNFESPVSTAT 60
Qy 61 AFAMLSLGTKAQTHDETLGLANFLNLTPEAQIHEGFOELRTLNOPDSQLQTONGLF 120
Db 61 AFAMLSLGTKAQTHDETLGLANFLNLTPEAQIHEGFOELRTLNOPDSQLQTONGLF 120
Qy 121 LSEGKLKVLDKFLEDVKLYHSRAFTVNFGDTEAKQKINDVYKGQTKIVDVLKELDRD 180
Db 121 LSEGKLKVLDKFLEDVKLYHSRAFTVNFGDTEAKQKINDVYKGQTKIVDVLKELDRD 180
Qy 181 TYPALVNYYFFKGKWERPFEVKDTEERDFHYDQVTTVKVPMKMRGLMENIOHCKKLSSWY 240
Db 181 TYPALVNYYFFKGKWERPFEVKDTEERDFHYDQVTTVKVPMKMRGLMENIOHCKKLSSWY 240
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Db 241 LSEGKLKVLDKFLEDVKLYHSRAFTVNFGDTEAKQKINDVYKGQTKIVDVLKELDRD 180
Qy 301 KSVLGQLGCTKVFNSNGADLSGYTEEAPLKLSKAVHAVITIDEKGTEAGAMFLERIPMS 360
Db 301 KSVLGQLGCTKVFNSNGADLSGYTEEAPLKLSKAVHAVITIDEKGTEAGAMFLERIPMS 360
Qy 361 IPEBVKPNKPFPVFLMBQNTKSPLFMGVNVNPTQKMSKSFKAQGCPPKSAQC 414
Db 361 IPEBVKPNKPFPVFLMBQNTKSPLFMGVNVNPTQKMSKSFKAQGCPPKSAQC 409

Qy 301 KSVLGQLGCTKVFNSNGADLSGYTEEAPLKLSKAVHAVITIDEKGTEAGAMFLERIPMS 360
Db 301 KSVLGQLGCTKVFNSNGADLSGYTEEAPLKLSKAVHAVITIDEKGTEAGAMFLERIPMS 360
Qy 361 IPEBVKPNKPFPVFLMBQNTKSPLFMGVNVNPTQKMSKSFKAQGCPPKSAQC 414

XX SQ sequence 580 AA;
XX Query Match 76.3%; Score 2040.5; DB 23; Length 580;
Best Local Similarity 96.6%; Pred. No. 3.1e-150;
Matches 400; Conservatory 2; Mismatches 7; Indels 5; Gaps 1;
Db 1 MEDPOGDAAQKTDPTSHHDQDHPTENKTPNLAEAFASLYROLAQSNTNFESPVSTAT 60
Db 1 MEDPOGDAAQKTDPTSHHDQDHPTENKTPNLAEAFASLYROLAQSNTNFESPVSTAT 60
Qy 61 AFAMLSLGTKAQTHDETLGLANFLNLTPEAQIHEGFOELRTLNOPDSQLQTONGLF 120
Db 61 AFAMLSLGTKAQTHDETLGLANFLNLTPEAQIHEGFOELRTLNOPDSQLQTONGLF 120
Qy 121 LSEGKLKVLDKFLEDVKLYHSRAFTVNFGDTEAKQKINDVYKGQTKIVDVLKELDRD 180
Db 121 LSEGKLKVLDKFLEDVKLYHSRAFTVNFGDTEAKQKINDVYKGQTKIVDVLKELDRD 180
Qy 181 TYPALVNYYFFKGKWERPFEVKDTEERDFHYDQVTTVKVPMKMRGLMENIOHCKKLSSWY 240
Db 181 TYPALVNYYFFKGKWERPFEVKDTEERDFHYDQVTTVKVPMKMRGLMENIOHCKKLSSWY 240
Qy 241 LLMKYLGNTATAIFFLPDECKLQHLENELTDITKFLENEDRRSASILHPLKSITGTYDL 300
Db 241 LLMKYLGNTATAIFFLPDECKLQHLENELTDITKFLENEDRRSASILHPLKSITGTYDL 300
Qy 301 KSVLGQLGCTKVFNSNGADLSGYTEEAPLKLSKAVHAVITIDEKGTEAGAMFLERIPMS 360
Db 301 KSVLGQLGCTKVFNSNGADLSGYTEEAPLKLSKAVHAVITIDEKGTEAGAMFLERIPMS 360
Qy 361 IPEBVKPNKPFPVFLMBQNTKSPLFMGVNVNPTQKMSKSFKAQGCPPKSAQC 414





Query	Match	Score	Length	DB	Length	DB
Best Local Similarity	76.1%	Score 2035;	DB 23;	Length 580;		
Matches	100.0%;	Pred. No.	8.3e-150;			
Matches	395; Conservative	0;	Mismatches	0;	Indels	0;
Matches						Gaps 0;
QY	1 MEDPQDAAQKTDPTSHHDQDHPTEPKITPNLAEFAFSLYRQLAHOSNSTNIFSPVSIAT	60				
DB	186 MEDPQDAAQKTDPTSHHDQDHPTEPKITPNLAEFAFSLYRQLAHOSNSTNIFSPVSIAT	245				
QY	61 AFAMSLGTRADTHDEILEGINFNITEIPPAQIHRGFQELRLTINQPSQLUTTGNGFL	120				
DB	246 AFAMSLGTRADTHDEILEGINFNITEIPPAQIHRGFQELRLTINQPSQLUTTGNGFL	305				
QY	121 ISEGGLKLVDKFLEDYKKLYSEAFTYEAKQINDYVEKGTOGKIVDVLVKELDRD	180				
DB	306 ISEGGLKLVDKFLEDYKKLYSEAFTYEAKQINDYVEGTQGKIVDVLVKELDRD	365				
QY	181 YFVALNYIFFKGKWRPFEVKDTEBEDFDHVQDQVTTVKVPMKRLGMFNHQCKKLSSWV	240				
DB	366 YFVALNYIFFKGKWRPFEVKDTEBEDFDHVQDQVTTVKVPMKRLGMFNHQCKKLSSWV	425				
QY	241 LIMKLYGNATAIFFLPDDEGKQHLHENELHDITKFLENEDRSASLHLPLKLSITGTIDL	300				
DB	426 LIMKLYGNATAIFFLPDDEGKQHLHENELHDITKFLENEDRSASLHLPLKLSITGTIDL	485				
QY	301 KSVLGQLGIFTKVFNSGADLSGVTEEAPLKSAYKHAVKAVLTIDEKGTEAAGMFLAIPMS	360				
DB	486 KSVLGQLGIFTKVFNSGADLSGVTEEAPLKSAYKHAVKAVLTIDEKGTEAAGMFLAIPMS	545				
QY	361 IPPEVKFNKPFPVFLMIEQNTKSPLFPMGKVYNTPTQK	395				
DB	546 IPPEVKFNKPFPVFLMIEQNTKSPLFPMGKVYNTPTQK	580				
RESULT 7						
ID	AAW59839	standard; Protein:	394 AA.			
XX						
AC	AAW59839;					
XX						
DT	20-NOV-1998	(first entry)				
XX						
DE	Nature protein sequence of alpha <sub>1</sub> -antitrypsin (AAT).					
XX						
Protein expression; monocytoid plant cell;						
KW	glycosylated alpha <sub>1</sub> -antitrypsin; AAT; glycosylated antithrombin III;					
KW	ATIII; human serum albumin; HSA; subtilisin BPN'; treatment; emphysema;					
KW	antithrombotic; blood replacement.					
XX						
OS	Homo sapiens.					
XX						
PN	W09836085-A1.					
XX						
PD	20-AUG-1998.					
XX						
PF	13-FEB-1998;	98WO-US03068.				
XX						
PR	13-FEB-1997;	97US-0038170.				
PR	13-FEB-1997;	97US-0038179.				
PR	13-FEB-1997;	97US-003818.				
PR	13-FEB-1997;	97US-003816.				
XX						
PA	(PHYT-) APPLIED PHYTOLOGICS INC.					
XX						
Rodriguez RL,	Sutliff TD;					
XX						
DR	WPI; 1998-467179/40.					
XX	N-PSDB; AAV41726.					
Expressing mature, glycosylated proteins in monocytoid plant cells - from chimeric genes including signal peptide sequence, specifically therapeutic agents and industrial enzymes						
XX						
Disclosure: Pages 28-29; 53pp; English.						
XX						
RESULT 8						
ID	AAU99873	standard; Protein:	394 AA.			
XX						
AC	AAU99873;					
XX						
DT	07-OCT-2002	(first entry)				
XX						
DE	Human alpha <sub>1</sub> -antitrypsin (AAT) protein.					
XX						
KW	Alpha-1-antitrypsin; AAT; human; protease inhibitor; malaria;					
KW	emphysema; asthma; chronic obstructive pulmonary disease; eczema;					
KW	cystic fibrosis; otitis media; otitis external; HIV; psoriasis;					
KW	human immunodeficiency virus; atop dermatitis; muscular dystrophy;					
KW	herpes; ulceration; sepsis; rheumatoid arthritis; periodontal disease;					
KW	tumour metastasis; tumour angiogenesis; osteoporosis; Paget's disease;					
KW	glomerulonephritis; scleroderma; Alzheimer's disease; hypertension.					
XX						
OS	Homo sapiens.					

PN	WO200250287-A2.	Qy	362 PPEVKFNKPKFVFLMIEONTKSPLFMGKVVNPTQK	395
XX	27-JUN-2002.	Db	361 PPEVKFNKPKFVFLMIEONTKSPLFMGKVVNPTQK	394
PD	XX			
XX	18-DEC-2001; 2001WO-US49256.			
PF	XX			
PR	18-DEC-2000; 2000US-256599P.			
PR	20-NOV-2001; 2001US-331366P.			
XX	PA (ARRI-) ARRIVA PHARM INC.			
XX	Barri PJ, Gibson HL, Pemberton P;			
PI	XX			
XX	WPI; 2002-500531/53.			
DR	N-PSDB; ABK80015.			
XX	Novel fusion protein useful for inhibiting protease activity associated with a disorder such as emphysema, asthma, comprises a first protease inhibitor comprising alpha 1-antitrypsin and a second protease inhibitor -			
PT	XX			
PT	Claim 25; Page 25-27; 134PP; English.			
XX	This invention relates to a novel fusion protein comprising a first protease inhibitor comprising an alpha-1-antitrypsin or its functionally active portion and a second protease inhibitor or its functionally active protein. The fusion proteins of the invention may act as an inhibitor of protease activity. The fusion protein of the invention is useful for inhibiting protease activity associated with a disorder such as emphysema, asthma, chronic obstructive pulmonary disease, cystic fibrosis, otitis media, otitis external or HIV infection, or for treating an individual suffering from or at risk for a disease or disorder involving unwanted protease activity. The proteins are useful for treating dermatological diseases such as atopic dermatitis, eczema and psoriasis, in inflammatory responses to viral infection, and for treating herpes infection, corneal or epidermal ulceration, chronic non-healing wounds, sepsis, rheumatoid arthritis, periodontal disease, tumour metastasis and tumour angiogenesis, gastric ulceration, osteoporosis, Paget's disease, glomerulonephritis, scleroderma, malaria, bacterial infection, Alzheimer's disease, hypertension and muscular dystrophy. The present sequence represents the human alpha-1-antitrypsin (AAT) protein used to create the fusion protein of the invention.			
XX	Sequence 394 AA;			
SQ				
Query Match	75.9%; Score 2010; DB 23; Length 394;			
Best Local Similarity	100.0%; Pred. No. 1.2e-149;			
Matches	394; Conservative 0; Mismatches 0; Indels 0; Gaps 0;			
Qy	2 EDPOGDAAGKTDTSHHDQDHTENKTKITPNLAEAFASLYRQLAHOSNSTNIFSPVSIATA	61		
Db	1 EDPOGDAAGKTDTSHHDQDHTENKTKITPNLAEAFASLYRQLAHOSNSTNIFSPVSIATA	60		
Qy	62 FAMSLIGTAKADTHDEILEGLNFNLTEIPQAQHFGQELRLTNOPDSQLQTGNGLF	121		
Db	61 FAMSLIGTAKADTHDEILEGLNFNLTEIPQAQHFGQELRLTNOPDSQLQTGNGLF	120		
Qy	122 SEGKLVDIFLEDYKKLYSEAFTVNFEDTEEAKQINDYERGTQKIVDVLKELDRDT	181		
Db	121 SEGKLVDIFLEDYKKLYSEAFTVNFEDTEEAKQINDYERGTQKIVDVLKELDRDT	180		
Qy	182 VFALVNYTFEKGKWERPFFKVDTEEEDPHDQVTTVKVPMMKRUGMENIQHCKKLSSWL	241		
Db	181 VFALVNYTFEKGKWERPFFKVDTEEEDPHDQVTTVKVPMMKRUGMENIQHCKKLSSWL	240		
Qy	242 LMKYLGNTAAIFFLPDEGLQHLENLTDITIKFLNEDRSASHLPLKLSITGTYDLK	301		
Db	241 LMKYLGNTAAIFFLPDEGLQHLENLTDITIKFLNEDRSASHLPLKLSITGTYDLK	300		
Qy	302 SVLGQLGITKVFNSGADISGVTEAPLKSKAVHKAVIDERGTGEAMPLEATPMSI	361		
Db	301 SVLGQLGITKVFNSGADISGVTEAPLKSKAVHKAVIDERGTGEAMPLEATPMSI	360		
Qy	182 VFALVNYIFFKGKWERPFFKVDQVTTVKVPMMKRUGMENIQHCKKLSSWL	241		

RESULT 9  
ID AAP40133 standard; Protein; 418 AA.  
XX  
AC AAP40133;  
XX DT 16-FEB-1992 (first entry)  
XX Sequence of human alpha-1-antitrypsin.  
XX Protease inhibitor; enzyme; proteolysis inhibitor; emphysema; therapy.  
XX DE Homo sapiens.  
XX FH Location/Qualifiers  
Peptide 1..24  
FT /label= signal  
Region 25..418  
PN EP103409-A.  
XX PD 21-MAR-1984.  
XX PF 12-AUG-1983; 83EP-0304668.  
XX PR 28-APR-1982; 83US-0489406.  
XX PR 13-AUG-1982; 82US-040809.  
XX PR 18-AUG-1982; 82US-040913.  
XX PR 01-JAN-1988; 88EP-0201179.  
XX PA (ZYMO-) ZYOMOS CORP.  
PA (BRIG-) BRIGHAM & WOMENS HO.  
PA (KAWA-) KAWASAKI.  
XX PI Kawasaki GH, Woodbury RG;  
XX DR N-PSDB; AAN40078.  
XX DR WPI: 1984-077108/13.  
XX DR N-PSDB; AAN40078.

Extra:chromosomal element for replication in yeast - with yeast promoter for regulation of glyolytic protein prodn.  
XX PS Disclosure: Fig 1A; 48pp; English.  
CC The inventors claim a DNA construct contg. a gene encoding human alpha-1-antitrypsin. A substantially pure, substantially unglycosylated mammalian alpha-1-antitrypsin is also claimed.  
XX SQ Sequence 418 AA;  
Query Match 75.9%; Score 2030; DB 5; Length 418;  
Best Local Similarity 100.0%; Fred. No. 1 3e-149;  
Matches 394; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Qy 2 EDPOGDAAGKTDTSHHDQDHTENKTKITPNLAEAFASLYRQLAHOSNSTNIFSPVSIATA 61  
Db 1 EDPOGDAAGKTDTSHHDQDHTENKTKITPNLAEAFASLYRQLAHOSNSTNIFSPVSIATA 60  
Qy 62 FAMSLIGTAKADTHDEILEGLNFNLTEIPQAQHFGQELRLTNOPDSQLQTGNGLF 121  
Db 61 FAMSLIGTAKADTHDEILEGLNFNLTEIPQAQHFGQELRLTNOPDSQLQTGNGLF 120  
Qy 122 SEGKLVDIFLEDYKKLYSEAFTVNFEDTEEAKQINDYERGTQKIVDVLKELDRDT 181  
Db 121 SEGKLVDIFLEDYKKLYSEAFTVNFEDTEEAKQINDYERGTQKIVDVLKELDRDT 180  
Qy 182 VFALVNYTFEKGKWERPFFKVDTEEEDPHDQVTTVKVPMMKRUGMENIQHCKKLSSWL 241  
Db 181 VFALVNYTFEKGKWERPFFKVDTEEEDPHDQVTTVKVPMMKRUGMENIQHCKKLSSWL 240  
Qy 242 LMKYLGNTAAIFFLPDEGLQHLENLTDITIKFLNEDRSASHLPLKLSITGTYDLK 301  
Db 241 LMKYLGNTAAIFFLPDEGLQHLENLTDITIKFLNEDRSASHLPLKLSITGTYDLK 300  
Qy 302 SVLGQLGITKVFNSGADISGVTEAPLKSKAVHKAVIDERGTGEAMPLEATPMSI 361  
Db 301 SVLGQLGITKVFNSGADISGVTEAPLKSKAVHKAVIDERGTGEAMPLEATPMSI 360  
Qy 182 VFALVNYIFFKGKWERPFFKVDQVTTVKVPMMKRUGMENIQHCKKLSSWL 241



(MERI-) MERISTEM THERAPEUTICS.

Gruber V, Olagnier B, Bournat P, Theisen M, Merot B;  
WPI; 1999-4699334/39.  
N-PSDB; AXX83518.

Production of alpha1-antitrypsin, and its variants, in cells of monocotyledonous plants, useful as serine protease inhibitors for therapy, e.g. of emphysema, in cosmetics and as reagents -

Claim 8; Fig 1; 67pp; French.

This sequence represents the coding region of the human alpha-1-antitrypsin (AT) gene. The invention relates to the production of AT in plant cells, especially monocotyledonous plants. Also produced are variants of the AT protein, in which the glycosylation pattern of the protein is altered. AT inhibits serine proteases, specifically neutrophil elastase (but also trypsin, cathepsin G, thrombin etc.) so protect pulmonary tissue against protease damage. AT are used to treat AT deficiency conditions, particularly pulmonary emphysema, cystic fibrosis, septic shock and rheumatism. The use of plants for the recombinant production of AT results in a product without risk of (sub)viral contamination. The recombinant AT had good activity and is stable, with low immunogenicity (associated with glycosylation patterns similar to the native Protein).

Alpha-1-antitrypsin from PDBA1.

Key Misc-difference	Location/Qualifiers	Value
	/note= "Met changed to Arg for alpha1AT-P; see CC"	
	GB2246779-A.	
	12-FEB-1992.	
	03-AUG-1990;	90GB-0017083.
	03-AUG-1990;	90GB-0017083.
	{DELT-} DELTA BIOTECH LTD.	
	Ballance DJ, Courtney MG;	
	WPI; 1992-051155/07.	
	N-PSDB; AAQ21125.	
	Antitumour molecules for treatment of neoplasms - comprises first region for binding to uPA receptor and second region for uPA inhibition	
	Disclosure: Fig 12; S70n; English	

A human alpha-haloperoxidase CDNA was modified to remove the 23 amino acid signal sequence and introduce a HindIII restriction site at the 3' end. The modified CDNA was cloned into M13mp19 to generate plasmid pDBA2 - see AAQ21123-24'. i.e. changing the codon for methionine 358 (337 in the sequence below) (ATG) such that it codes for arginine (AGG). See also AAQ21119 and AAQ21121-25.							
Sequence	393 AA;						
Very Match	75.6%	Score	2022:	DB 1.3;	Length	393;	
st. Local Similarity	99.7%	pred. No.	5e-149;				
Matches	392;	Mismatches	1;				
		Indels	0;				
		Ga					
3	DPGDAAQKTDPSHHDQDHPFTNKKITPNLAEAFASLYROLAHQSNTNSTNFFSPVSIATAF						
1	DEQGDAAQKTDPSHHDQDHPFTNKKITPNLAEAFASLYROLAHQSNTNSTNFFSPVSIATAF						
63	AMLSLGTKADTDEILBLNFIETIPPAQIHEGFOELLRTLNQPDSQLQTGTGNGFLS						
61	AMLSLGTKADTDEILBLNFIETIPPAQIHEGFOELLRTLNQPDSQLQTGTGNGFLS						
123	BGLKLVDKFLEVKKLYHSEATIVNFGDTEEAKQINIDYVEKGQKIVDVLVKELDRDTV						
121	BGLKLVDKFLEVKKLYHSEATIVNFGDTEEAKQINIDYVEKGQKIVDVLVKELDRDTV						
183	FALVNVTFFKGKWERPFPVYKDTEEDPHVDQTTVYKVPMMKRLGMENIQHKCKLSSWVLL						
181	FALVNVTFFKGKWERPFPVYKDTEEDPHVDQTTVYKVPMMKRLGMENIQHKCKLSSWVLL						
243	MKYLGNATAIFFLPDESKLQHLENELTHDIKFLENEDRASSLHLPKLSTGTYDLKS						
241	MKYLGNATAIFFLPDESKLQHLENELTHDIKFLENEDRASSLHLPKLSTGTYDLKS						
303	VLGQLGITKVPSGADLSGVTEAPLKSKAVHKAVLTIDEKGTEAAGMFLERIPMSIP						
301	VLGQLGITKVPSGADLSGVTEAPLKSKAVHKAVLTIDEKGTEAAGMFLERIPMSIP						
363	PEVKENKPKFVFLMIEONTKSPFLMGKVYNPTOK	395					
361	PDVKENKPKFVFLMIEONTKSPFLMGKVYNPTOK	393					

RESULT 13  
AAR71969

AAR71969;	QY	302	SVLQLGLTGVKVSNGADLSGTYEEAPLKSLKAVHKAVLTTIDEKGTEAAAGMFLAIPMSI	361
X AAR71969;	Db	325	SVLQLGLTGVKVSNGADLSGTYEEAPLKSLKAVHKAVLTTIDEKGTEAAAGMFLAIPMSI	384
X 18-OCT-1995 (first entry)	QY	362	PPEVKENPFPVFVFLMIEQNTKSPLFMGVNPQQ 418	
X Human alpha-1-trypsin.	Db	385	RPEVKENKRPFPVFVFLMIEQNTKSPLFMGVNPQQ 418	
X Alpha-1-trypsin; protease-inhibitor.				
X Homo sapiens.				
X				
Key	Location/Qualifiers			
Peptide	1..24			
	/label= Sig_peptide			
X				
US399684-A.				
X				
21-MAR-1995.				
DD				
X				
20-MAY-1982;	82US-0380310.			
XX				
20-MAY-1982;	82US-0380310.			
XX				
07-FEB-1984;	84US-0638980.			
XX				
03-MAR-1987;	87US-002243.			
XX				
15-DEC-1987;	87US-0133193.			
XX				
16-SEP-1988;	88US-0246912.			
XX				
22-AUG-1989;	89US-0398288.			
XX				
11-MAR-1991;	91US-0666450.			
XX				
18-NOV-1992;	92US-0979556.			
XX				
02-JUL-1993;	93US-0086142.			
XX				
DR				
AK089254.				
(WASH-) WASHINGTON RES FOUND.				
PA				
Davie EW, Kurachi K, Thirumalachary C, Woo SLC;				
X				
Human alpha1-antitrypsin (alpha1-AT) cDNA sequence - can be used for				
the expression of alpha1-AT.				
X				
Disclosure: Fig.1; 15pp; English.				
X				
The sequence of human alpha-1-antitrypsin encoded by an isolated				
cDNA clone is given in AAC1969. Expression of the cDNA in host cell				
transformants allowed production of recombinant alpha-1-antitrypsin.				
X				
Sequence 418 AA;				
SQ				
Query Match	75.6%	Score 2021;	DB 16;	Length 418;
Best Local Similarity	99.7%	Pred. No.	6.5e-149;	
Matches	393;	Conservative	0; Mismatches	0;
QY	2	EDPGDAAQKDTSHHDQDHPTENKTPNLEFAFSLYROLAHOSNSTNFSPSVAATA	61	
Db	25	EDPGDAAQKDTSHHDQDHPTENKTPNLEFAFSLYROLAHOSNSTNFSPSVAATA	84	
QY	62	FAMLSLTGKTDTHDETLGFLNLTEPAQIHEGQELLFTLNQPSDQLQTGNGLFL	121	
Ddb	85	FAMLSLTGKTDTHDETLGFLNLTEPAQIHEGQELLFTLNQPSDQLQTGNGLFL	144	
QY	122	SEGKLKYDKFELDKVLYHSAFTVNGFDTEBAKKQINDYVEKGTKIVDLVKELDRDT	181	
Ddb	145	SEGKLKYDKFELDKVLYHSAFTVNGFDTEBAKKQINDYVEKGTKIVDLVKELDRDT	204	
QY	162	VEFLVNYIFFGKWERPFEEVKYOTEEEDFHVDQVTTVYKPMKRLGMFNIOCKKLSWSWLL	241	
QY	205	VFLVNYIFFGKWERPFEEVKYOTEEEDFHVDQVTTVYKPMKRLGMFNIOCKKLSWSWLL	264	
Ddb	242	LMKYLNATAIFLPDGSKLQLHENELTHDITKFLNEEDRSASLHLPKLSITGTYDLK	301	
QY	265	LMKYLNATAIFLPDGSKLQLHENELTHDITKFLNEEDRSASLHLPKLSITGTYDLK	324	
QY	122	SEGLKLKYDKFELDKVLYHSAFTVNGFDTEBAKKQINDYVEKGTKIVDLVKELDRDT	181	
QY	162	FAMLSLTGKTDTHDETLGFLNLTEPAQIHEGQELLFTLNQPSDQLQTGNGLFL	144	
QY	205	FAMLSLTGKTDTHDETLGFLNLTEPAQIHEGQELLFTLNQPSDQLQTGNGLFL	121	
QY	25	FAMLSLTGKTDTHDETLGFLNLTEPAQIHEGQELLFTLNQPSDQLQTGNGLFL	144	
QY	62	FAMLSLTGKTDTHDETLGFLNLTEPAQIHEGQELLFTLNQPSDQLQTGNGLFL	144	
QY	85	FAMLSLTGKTDTHDETLGFLNLTEPAQIHEGQELLFTLNQPSDQLQTGNGLFL	144	
QY	122	SEGLKLKYDKFELDKVLYHSAFTVNGFDTEBAKKQINDYVEKGTKIVDLVKELDRDT	181	

RESULT 15  
 AAY78890 ID AAY78890 standard; Protein: 418 AA.  
 XX AC  
 XX DT 19-MAY-2000 (first entry)  
 XX DE Human alpha-antitrypsin amino acid sequence.  
 XX KW Alpha-antitrypsin; neutrophil elastase inhibitor; human;  
 KW chronic obstructive pulmonary emphysema; infantile liver cirrhosis.  
 XX OS Homo sapiens.  
 XX PN US6025161 A.  
 XX PD 15-FEB-2000.  
 XX PF 20-JAN-1998; 98US-0009581.  
 PR 07-JUN-1995; 95US-0479545.  
 PR 20-MAY-1987; 82US-0380810.  
 PR 07-FEB-1984; 84US-0638980.  
 PR 03-MAR-1987; 87US-0022543.  
 PR 15-DEC-1987; 87US-0131390.  
 PR 16-SEP-1988; 88US-0246912.  
 PR 22-AUG-1989; 89US-0399288.  
 PR 11-MAR-1991; 91US-0666450.  
 PR 18-NOV-1992; 92US-097556.  
 PR 02-JUL-1993; 93US-0086642.  
 XX PA (WASH-) WASHINGTON RES FOUND.  
 XX PI Woo SLC, Thirumalachary C, Kurachi K, Davie EW;  
 XX DR WPI: 2000-181811/16.  
 XX DR N-PSDB; AAZ9099.

for the production of alpha1-antitrypsin.

CC XX sequence 418 AA;  
 SQ Query Match 75.6%; Score 2021; DB 21;  
 Best Local Similarity 99.7%; Pred. No. 6.5e-149;  
 Matches 393; Conservative 0; Mismatches 1; Indels 0; Gaps 0;  
 Qy 2 EDPGDAAQKRTDSHHDQHPTNKITNLAEAFSRYROLAQHSNTNIFSPVSDATA 61  
 Db 25 EDPGDAAQKTDSSHDDQHPTNKITNLAEAFSRYROLAQHSNTNIFSPVSDATA 84  
 Qy 62 FAMLSLGTKADTHDEILEGLDNFLTEPEAQIHEGFDELLRTLNQPSOLQTGNGFL 121  
 Db 85 FAMLSLGTKADTHDEILEGLDNFLTEPEAQIHEGFDELLRTLNQPSOLQTGNGFL 144  
 Qy 122 SEGKLVDKFKLEDYKKLYHSEATVNFQDTEAKKOINDYVEKGTOCKIVDVLVKELDRDT 181  
 Db 145 SEGKLVDKFKLEDYKKLYHSEATVNFQDTEAKKOINDYVEKGTOCKIVDVLVKELDRDT 204  
 Qy 182 VFAVLNVYIFFKGWPERPEVKDTEEDFHVDQVTYVPMKRLGMENIQCKKLSWVL 241  
 Db 205 VFAVLNVYIFFKGWPERPEVKDTEEDFHVDQVTYVPMKRLGMENIQCKKLSWVL 264  
 Qy 242 LMKYLGNTAIAFLPDEGKQLHLENELTHDITTKFLNEQDRRSASLHLPKLSITGTYDLK 301  
 Db 265 LMKYLGNTAIAFLPDEGKQLHLENELTHDITTKFLNEQDRRSASLHLPKLSITGTYDLK 324  
 Qy 302 SVGQLGTTKVFSGNADLSGVTEAPLKLASKAVHKAVLTIDEKGTEAGAMFLEALPMSI 361  
 Db 325 SVGQLGTTKVFSGNADLSGVTEAPLKLASKAVHKAVLTIDEKGTEAGAMFLEALPMSI 384  
 Qy 362 PPEVKENKPFVFLMIEQNTKSPLFMGVVNPTQK 395  
 Db 385 RPEVKENKPFVFLMIEQNTKSPLFMGVVNPTQK 418

Search completed: November 30, 2002, 12:35:00  
 Job time : 29 secs

Prepar ing alpha1-antitrypsin for inhibiting neutrophil elastase  
 involves transfecting host cell with vector comprising  
 alpha1-antitrypsin DNA sequence that hybridizes to human  
 alpha1-antitrypsin cDNA, or its complement -  
 Claim 1: Fig 1; 16pp; English.  
 CC This sequence represents the human alpha1-antitrypsin amino acid  
 sequence. Alpha1-antitrypsin is an important protease inhibitor, the  
 major function of which is to inhibit neutrophil elastase; low levels of  
 alpha1-antitrypsin in the blood are associated with chronic obstructive  
 pulmonary emphysema and infantile liver cirrhosis. A vector comprising a  
 mammalian alpha1-antitrypsin DNA sequence that hybridizes to human  
 alpha1-antitrypsin cDNA can be introduced into a host cell in a method

